(19) World Intellectual Property Organization

International Bureau



(43) International Publication Date 24 June 2004 (24.06.2004)

PCT

(10) International Publication Number WO 2004/053089 A3

(51) International Patent Classification⁷: C07K 1/00, C12Q 1/68, 1/34, C12P 19/34, C07H 21/02

(21) International Application Number:

PCT/US2003/039087

(22) International Filing Date:

10 December 2003 (10.12.2003)

(25) Filing Language:

English

(26) Publication Language:

English

(30) Priority Data:

60/432,432

11 December 2002 (11.12.2002) US

- (71) Applicant (for all designated States except US): THE RESEARCH FOUNDATION OF STATE UNIVERSITY OF NEW YORK [US/US]; P.O. Box 9, Albany, NY 12201-0009 (US).
- (72) Inventors; and
- (75) Inventors/Applicants (for US only): MCALLISTER, William, T. [US/US]; 196 Water Street, Perth Amboy, NJ 13084 (US). KUKARIN, Alexander [RU/US]; 6811 Burus, E3, Forest Hills, NY 11375 (US).
- (74) Agent: HOFFMAN, Michael, F.; Hoffman, Warnick & D'Alessandro LLC, Three E-Comm Square, Albany, NY 12207 (US).

- (81) Designated States (national): AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW.
- (84) Designated States (regional): ARIPO patent (BW, GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW), Eurasian patent (AM, AZ, BY, KG, KZ, MD, RU, TJ, TM), European patent (AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK, TR), OAPI patent (BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG).

Declaration under Rule 4.17:

of inventorship (Rule 4.17(iv)) for US only

Published:

- with international search report
- before the expiration of the time limit for amending the claims and to be republished in the event of receipt of amendments
- (88) Date of publication of the international search report: 16 September 2004

For two-letter codes and other abbreviations, refer to the "Guidance Notes on Codes and Abbreviations" appearing at the beginning of each regular issue of the PCT Gazette.

(54) Title: IMPROVED METHODS AND MATERIALS FOR REDUCING PRODUCTION OF ABERRANT PRODUCTS DURING RNA SYNTHESIS

(57) Abstract: The mutant T7, T3 and SP6 RNA polymerase enzymes have a number of novel properties that would be desirable for those who wish to use the phage RNAP to synthesize homogeneous RNA products. These includes: reduced synthesis of aberrant products on templates that have protruding 3' ends in the non-template strand, decreased addition of a non-templated nucleotide to the 37 end of transcript; increased yields of products on templates that termioate in a G:c-rich context. The mutations include deletions at positions 172 and 173 of the T7 RNA polymerase, deletions at positions 140-143 of the SP6 RNA polymerase and deletions at positions 173 and 174 of the T3 RNA polymerase.





INTERNATIONAL SEARCH REPORT

International application No.

PCT/US03/39087

A. CLASSIFICATION OF SUBJECT MATTER IPC(7) : C07K 1/00; C12Q 1/68, 1/34; C12P 19/34; C07H 21/02 US CL : 530/350; 435/6, 18, 91.2; 536/23.1				
According to International Patent Classification (IPC) or to both national classification and IPC B. FIELDS SEARCHED				
Minimum documentation searched (classification system followed by classification symbols) U.S.: 530/350; 435/6, 18, 91.2; 536/23.1				
Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched				
Electronic data base consulted during the international search (name of data base and, where practicable, search terms used) Please See Continuation Sheet				
C. DOCUMENTS CONSIDERED TO BE RELEVANT				
Category *	Citation of document, with indication, where ap		Relevant to claim No.	
ĸ	LYAKHOV et al. Mutant bacteriophage T7 RNA pe		1-6, 8-14	
Ÿ	properties. J. Mol. Biol. 1997, Vol. 269, pages 28-	40, see entire document.	7, 15	
x	GOPAL et al. Characterization of structural feature	s important for T7 RNAP elongation	1-5, 8-13	
	complex stability reveals competing complex confor	rmations and a role for the non-		
Y	template strand in RNA displacement. J. Mol. Biol. entire document.	. 1999, Vol 290, pages 411-431, see	6, 7, 14, 15	
X, P	EP 1 403 364 A1 (NIPPON GENETECH CO. LTD	0.) 19 December 2002 (19.12.2002),	1-15	
Y,P	page 7, lines 5-55. US 6,524,828 B1 (LIAO et al) 25 February 2003 (2	25 02 2003) see entire document.	1-15	
			·	
Further	documents are listed in the continuation of Box C.	See patent family annex.	4.5	
* S	pecial categories of cited documents:	"T" later document published after the inte date and not in conflict with the applic	emational filing date or priority	
	defining the general state of the art which is not considered to be	principle or theory underlying the inve	ention	
•	plication or patent published on or after the international filing date	"X" document of particular relevance; the considered novel or cannot be conside when the document is taken alone	claimed invention cannot be cred to involve an inventive step	
"L" document establish specified)	which may throw doubts on priority claim(s) or which is cited to the publication date of another citation or other special reason (as	"Y" document of particular relevance; the considered to involve an inventive step combined with one or more other sucl	p when the document is	
"O" document	referring to an oral disclosure, use, exhibition or other means	being obvious to a person skilled in th		
"P" document published prior to the international filing date but later than the "&" document member of the same patent family priority date claimed		family		
	ctual completion of the international search	Date of mailing of the international sea	Ø4 AUG 2004	
Name and mailing address of the ISA/US Mail Stop PCT, Atm: ISA/US Commissioner for Patents P.O. Box 1450 Alexandria, Virginia 22313-1450 Facsimile No. (703) 305-3230		Jeffley Fredman Telephone No. 703-308-0196		
1 acamine IAC	. (100) 300-3200			

	PCT/US03/39087			
INTERNATIONAL SEARCH REPORT	101/000/07-07			
MATERIALISM DESIRON MET OFF				
	· ·			
·				
•				
Continuation of B. FIELDS SEARCHED Item 3:				
EAST, DERWENT, MEDLINE, BIOSIS, CAPLUS search terms: RNA, polymerase, mutants, variants, polymorphisms, alleles, T7, SP6, T3, 173, 174, 172, 140, 143, deletion.				
4 11 41-1 7	27 cpc 72 173 174 172 140 143 deletion			
search terms: RNA, polymerase, mutants, variants, polymorphisms, alleles, T	7, SP6, T3, 173, 174, 172, 140, 143, deletion.			
search terms: RNA, polymerase, mutants, variants, polymorphisms, alleles, T	7, SP6, T3, 173, 174, 172, 140, 143, deletion.			
search terms: RNA, polymerase, mutants, variants, polymorphisms, alleles, T	7, SP6, T3, 173, 174, 172, 140, 143, deletion.			
search terms: RNA, polymerase, mutants, variants, polymorphisms, alleles, T	7, SP6, T3, 173, 174, 172, 140, 143, deletion.			
search terms: RNA, polymerase, mutants, variants, polymorphisms, alleles, T	7, SP6, T3, 173, 174, 172, 140, 143, deletion.			
search terms: RNA, polymerase, mutants, variants, polymorphisms, alleles, T	7, SP6, T3, 173, 174, 172, 140, 143, deletion.			
search terms: RNA, polymerase, mutants, variants, polymorphisms, alleles, T	7, SP6, T3, 173, 174, 172, 140, 143, deletion.			
search terms: RNA, polymerase, mutants, variants, polymorphisms, alleles, T	7, SP6, T3, 173, 174, 172, 140, 143, deletion.			
search terms: RNA, polymerase, mutants, variants, polymorphisms, alleles, T	7, SP6, T3, 173, 174, 172, 140, 143, deletion.			
search terms: RNA, polymerase, mutants, variants, polymorphisms, alleles, T	7, SP6, T3, 173, 174, 172, 140, 143, deletion.			
search terms: RNA, polymerase, mutants, variants, polymorphisms, alleles, T	7, SP6, T3, 173, 174, 172, 140, 143, deletion.			
search terms: RNA, polymerase, mutants, variants, polymorphisms, alleles, T	7, SP6, T3, 173, 174, 172, 140, 143, deletion.			
	7, SP6, T3, 173, 174, 172, 140, 143, deletion.			
	7, SP6, T3, 173, 174, 172, 140, 143, deletion.			
	7, SP6, T3, 173, 174, 172, 140, 143, deletion.			
	7, SP6, T3, 173, 174, 172, 140, 143, deletion.			
	7, SP6, T3, 173, 174, 172, 140, 143, deletion.			
	7, SP6, T3, 173, 174, 172, 140, 143, deletion.			
	7, SP6, T3, 173, 174, 172, 140, 143, deletion.			

Form PCT/ISA/210 (second sheet) (July 1998)